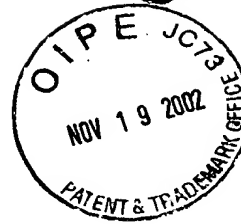


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<120> METHODS AND COMPOUNDS FOR MODULATING NUCLEAR RECEPTOR
COACTIVAOR BINDING

<130> 9811-008-999

<140> 09/281,717

<141> 1999-03-30

<150> US 60/079,956

<151> 1998-03-30

<160> 60

<170> PatentIn version 3.0

<210> 1

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<212> PRT

<213> Homo sapiens

<220>

<221> Variant

<222> (2)..(3)

<223> Xaa is any amino acid

<400> 1

Leu Xaa Xaa Leu Leu
1 5

<210> 2

<211> 6

<212> PRT

<213> Homo sapiens

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<221> Variant

<222> (3)..(4)

<223> Xaa is any amino acid

<400> 2

Ile Leu Xaa Xaa Leu Leu
1 5

<210> 3

<211> 5

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<222> (2)..(3)

<223> Xaa is any amino acid

<400> 3

Phe Xaa Xaa Leu Trp
1 5

<210> 4

<211> 5

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<220>

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<222> (2)..(3)

<223> Xaa is any amino acid

<400> 4

Phe Xaa Xaa Ala Leu
1 5

<210> 5

<211> 34

<212> PRT

<213> Homo sapiens

<400> 5

Ala Glu Gly His Ser Arg Leu His Asp Ser Lys Gly Gln Thr Lys Leu
1 5 10 15

Leu Gln Leu Leu Thr Thr Lys Ser Glu Gln Met Glu Pro Ser Pro Leu
20 25 30

Ala Ser

<210> 6

<211> 34

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<213> Homo sapiens

<220>

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<222> (15)..(15)

<223> Ile --> Ala

<220>

<221> MUTAGEN

<222> (16)..(16)

<223> Leu --> Ala

<220>

<221> MUTAGEN

<222> (19)..(20)

<223> Leu --> Ala

<220>

<221> MUTAGEN

<222> (16)..(20)

<223> Leu(16) --> Ala; Leu (20) -->Ala

<220>

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<222> (15)..(16)

<223> IleLeu --> AlaAla

<220>

<221> MUTAGEN

<222> (17)..(18)

<223> HisArg -->AlaAla

<220>

<221> MUTAGEN

<222> (15)..(15)

<223> Ile -->Phe

<220>

<221> MUTAGEN

<222> (16)..(16)

<223> Leu -->Phe

<220>

<221> MUTAGEN

<222> (19) .. (19)

<223> Leu -->Phe

<220>

<221> MUTAGEN

<222> (20) .. (20)

<223> Leu -->Phe

<400> 6

Pro Gly Ser Thr His Gly Thr Ser Leu Lys Glu Lys His Lys Ile Leu
1 5 10 15

His Arg Leu Leu Gln Asp Ser Ser Ser Pro Val Asp Leu Ala Lys Leu
20 25 30

Thr Ala

<210> 7

<211> 31

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<213> Homo sapiens

<400> 7

Glu Pro Ala Ser Pro Lys Lys Lys Glu Asn Ala Leu Leu Arg Tyr Leu
1 5 10 15

Leu Asp Lys Asp Asp Thr Lys Asp Ile Gly Leu Pro Glu Ile Thr
20 25 30

<210> 8

<211> 34

<212> PRT

<213> Homo sapiens

<400> 8

Ala Asp Gly Gln Ser Arg Leu His Asp Ser Lys Gly Gln Thr Lys Leu
1 5 10 15

Leu Gln Leu Leu Thr Thr Lys Ser Glu Gln Met Glu Pro Ser Pro Leu
20 25 30

Ala Ser

<210> 9

<211> 34

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<213> Homo sapiens

<400> 9

Ser Gly Ser Thr His Gly Thr Ser Leu Lys Glu Lys His Lys Ile Leu
1 5 10 15

His Arg Leu Leu Gln Asp Ser Ser Ser Pro Val Asp Leu Ala Lys Leu
20 25 30

Thr Ala

<210> 10

<211> 31

<212> PRT

<213> Homo sapiens

<400> 10

Glu Pro Val Ser Pro Lys Lys Lys Glu Asn Ala Leu Leu Arg Tyr Leu
1 5 10 15

Leu Asp Lys Asp Asp Thr Lys Asp Ile Gly Leu Pro Glu Ile Thr
20 25 30

<210> 11

<211> 34

<212> PRT

<213> Homo sapiens

<400> 11

Ala Glu Gly His Ser Arg Leu His Asp Ser Lys Gly Gln Thr Lys Leu
1 5 10 15

Leu Gln Leu Leu Thr Thr Lys Ser Glu Gln Met Glu Pro Ser Pro Leu
20 25 30

Pro Ser

<210> 12

<211> 34

<212> PRT

<213> Homo sapiens

<400> 12

Pro Gly Ser Thr His Gly Thr Ser Leu Lys Glu Lys His Lys Ile Leu
1 5 10 15

His Arg Leu Leu Gln Asp Ser Ser Ser Pro Val Asp Leu Ala Lys Leu
20 25 30

Thr Ala

<210> 13

<211> 31

<212> PRT

<213> Homo sapiens

<400> 13

Glu Pro Ala Ser Pro Lys Lys Lys Glu Asn Ala Leu Leu Arg Tyr Leu
1 5 10 15

Leu Asp Lys Asp Asp Thr Lys Asp Ile Gly Leu Pro Ser Ile Thr
20 25 30

<210> 14

<211> 34

<212> PRT

<213> Homo sapiens

<400> 14

Ala Glu Asn Gln Arg Gly Pro Leu Glu Ser Lys Gly His Lys Lys Leu
1 5 10 15

Leu Gln Leu Leu Thr Cys Ser Ser Glu Asp Arg Gly His Ser Ser Leu
20 25 30

Thr Asn

<210> 15

<211> 34

<212> PRT

<213> Homo sapiens

<400> 15

Thr	Ser	Asn	Met	His	Gly	Ser	Leu	Leu	Gln	Glu	Lys	His	Arg	Ile	Leu
1				5					10					15	

His	Lys	Leu	Leu	Gln	Asn	Gly	Asn	Ser	Pro	Ala	Glu	Val	Ala	Lys	Ile
			20					25					30		

Thr Ala

<210> 16

<211> 32

<212> PRT

<213> Homo sapiens

<400> 16

Glu	Gln	Leu	Ser	Pro	Lys	Lys	Lys	Glu	Asn	Asn	Ala	Leu	Leu	Arg	Tyr
1				5				10						15	

Leu	Leu	Asp	Arg	Asp	Asp	Pro	Ser	Asp	Val	Leu	Ala	Lys	Lys	Leu	Gln
			20					25					30		

<210> 17

<211> 34

<212> PRT

<213> Homo sapiens

<400> 17

Ala	Glu	Asn	Gln	Arg	Gly	Pro	Leu	Glu	Ser	Lys	Gly	His	Lys	Lys	Leu
1				5				10						15	

Leu	Gln	Leu	Leu	Thr	Cys	Ser	Ser	Asp	Asp	Arg	Gly	His	Ser	Ser	Leu
			20					25					30		

Thr Asn

<210> 18

<211> 34

<212> PRT

<213> Homo sapiens

<400> 18

Thr Ser Asn Met His Gly Ser Leu Leu Gln Glu Lys His Arg Ile Leu
1 5 10 15

His Lys Leu Leu Gln Asn Gly Asn Ser Pro Ala Glu Val Ala Lys Ile
20 25 30

Thr Ala

<210> 19

<211> 32

<212> PRT

<213> Homo sapiens

<400> 19

Glu Gln Leu Ser Pro Lys Lys Lys Glu Asn Asn Ala Leu Leu Arg Tyr
1 5 10 15

Leu Leu Asp Arg Asp Asp Pro Ser Asp Ala Leu Ser Lys Glu Leu Gln
20 25 30

<210> 20

<211> 34

<212> PRT

<213> Homo sapiens

<400> 20

Ser Glu Thr Pro Arg Gly Pro Leu Glu Ser Lys Gly His Lys Lys Leu
1 5 10 15

Leu Gln Leu Leu Thr Cys Ser Ser Glu Asp Arg Gly His Ser Ser Leu
20 25 30

Thr Asn

<210> 21

<211> 34

<212> PRT

<213> Homo sapiens

<400> 21

Thr Ser Asn Val His Gly Ser Leu Leu Gln Glu Lys His Arg Ile Leu
1 5 10 15

His Lys Leu Leu Gln Asn Gly Asn Ser Pro Ala Glu Val Ala Lys Ile
20 25 30

Thr Ala

<210> 22

<211> 32

<212> PRT

<213> Homo sapiens

<400> 22

Glu Gln Leu Ser Pro Lys Lys Lys Glu Asn Asn Ala Leu Leu Arg Tyr
1 5 10 15

Leu Leu Asp Arg Asp Asp Pro Ser Asp Ala Leu Ser Lys Glu Leu Gln
20 25 30

<210> 23

<211> 32

<212> PRT

<213> Homo sapiens

<400> 23

Ser Glu Gly Asp Ser Lys Tyr Ser Gln Thr Ser His Lys Leu Val Gln
1 5 10 15

Leu Leu Thr Thr Thr Ala Glu Gln Gln Leu Arg His Ala Asp Ile Asp
20 25 30

<210> 24

<211> 33

<212> PRT

<213> Homo sapiens

<400> 24

Thr Cys Pro Ser Ser His Ser Ser Leu Thr Glu Arg His Lys Ile Leu
1 5 10 15

His Arg Leu Leu Gln Glu Gly Ser Pro Ser Asp Ile Thr Thr Leu Ser
20 25 30

Val

<210> 25

<211> 34

<212> PRT

<213> Homo sapiens

<400> 25

Glu	Leu	Asp	Ala	Ala	Lys	Lys	Lys	Glu	Ser	Lys	Asp	His	Gln	Leu	Leu
1				5				10						15	
Arg	Tyr	Leu	Leu	Asp	Lys	Asp	Glu	Lys	Asp	Leu	Arg	Ser	Thr	Pro	Asn
			20					25					30		

Leu Cys

<210> 26

<211> 34

<212> PRT

<213> Homo sapiens

<220>

<221> SITE

<222> (2)..(2)

<223> Xaa is a negatively charged amino acid

<220>

<221> SITE

<222> (17)..(17)

<223> Xaa is a hydrophobic amino acid

<220>

<221> SITE

<222> (25)..(25)

<223> Xaa is a negatively charged amino acid

<220>

<221> Variant

<222> (1)..(1)

<223> Xaa is any amino acid

<220>

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<222> (3)..(9)
<223> Xaa is any amino acid

<220>
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<222> (11)..(14)
<223> Xaa is any amino acid

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<222> (22)..(24)
<223> Xaa is any amino acid

<220>
<221> Variant
<222> (26)..(34)
<223> Xaa is any amino acid

<400> 26
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Ser Xaa Xaa Xaa Xaa Lys Leu
1 5 10 15
Xaa Gln Leu Leu Thr Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa
20 25 30
Xaa Xaa

<210> 27
<211> 34
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<221> SITE
<222> (12)..(12)

<223> Xaa is a positively charged amino acid

<220>

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<222> (14)..(14)

<223> Xaa is a positively charged amino acid

<220>

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<222> (18)..(18)

<223> Xaa is a positively charged amino acid

<220>

<221> SITE

<222> (28)..(28)

<223> Xaa is a negatively charged amino acid

<220>

<221> SITE

<222> (29)..(29)

<223> Xaa is a hydrophobic amino acid

<220>

<221> SITE

<222> (32)..(32)

<223> Xaa is a hydrophobic amino acid

<400> 27

Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Leu	Xaa	Glu	Xaa	His	Xaa	Ile	Leu
1				5				10						15	

His	Xaa	Leu	Leu	Gln	Xaa	Xaa	Xaa	Ser	Pro	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa
		20						25					30		

Xaa Xaa

<210> 28

<211> 34
<212> PRT
<213> Homo sapiens

<220>
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<222> (22)..(22)
<223> Xaa is a positively charged amino acid

<220>
<221> SITE
<222> (24)..(24)
<223> Xaa is a negatively charged amino acid

<220>
<221> SITE
<222> (33)..(33)
<223> Xaa is a hydrophobic amino acid

<220>
<221> Variant
<222> (2)..(5)
<223> Xaa is any amino acid

<220>
<221> Variant
<222> (10)..(14)
<223> Xaa is any amino acid

<220>
<221> Variant
<222> (25)..(32)
<223> Xaa is any amino acid

<220>

<221> Variant

<222> (34)..(34)

<223> Xaa is any amino acid

<400> 28

Glu	Xaa	Xaa	Xaa	Xaa	Lys	Lys	Lys	Glu	Xaa	Xaa	Xaa	Xaa	Xaa	Leu	Leu
1				5					10					15	

Arg	Tyr	Leu	Leu	Asp	Xaa	Asp	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa	Xaa
			20				25						30		

Xaa Xaa

<210> 29

<211> 18

<212> PRT

<213> Homo sapiens

<400> 29

Thr	Ser	Leu	Lys	Glu	Lys	His	Lys	Leu	Leu	Arg	Tyr	Leu	Leu	Gln	Asp
1				5					10					15	

Ser Ser

<210> 30

<211> 33

<212> PRT

<213> Homo sapiens

<220>

<221> MUTAGEN

<222> (5)..(5)

<223> Thr --> Arg (T281R)

<220>

<221> MUTAGEN

<222> (8)..(8)

<223> Val --> Arg (V284R)

<220>

<221> MUTAGEN

<222> (9)..(9)

<223> Asp --> Ala (D285A)

<220>

<221> MUTAGEN

<222> (12)..(12)

<223> Lys --> Ala (K288A)

<220>

<221> MUTAGEN

<222> (22)..(22)

<223> Cys --> Arg (C298R)

<220>

<221> MUTAGEN

<222> (26)..(26)

<223> Ile --> Arg (I302R)

<220>

<221> MUTAGEN

<222> (30)..(30)

<223> Lys --> Ala (K306A)

<400> 30

Thr	Pro	Ala	Ile	Thr	Arg	Val	Val	Asp	Phe	Ala	Lys	Lys	Leu	Pro	Met
1				5					10					15	

Phe	Cys	Glu	Leu	Pro	Cys	Glu	Asp	Gln	Ile	Ile	Leu	Leu	Lys	Gly	Cys
			20					25					30		

Cys

<210> 31
<211> 12
<212> PRT
<213> Homo sapiens

<220>
<221> MUTAGEN
<222> (5)..(5)
<223> Leu --> Arg (L454R)

<220>
<221> MUTAGEN
<222> (7)..(7)
<223> Leu --> Arg (L456R)

<220>
<221> MUTAGEN
<222> (8)..(8)
<223> Glu -->Lys (E457K)

<400> 31
Leu Phe Pro Pro Leu Phe Leu Glu Val Phe Glu Asp
1 5 10

<210> 32
<211> 33
<212> PRT
<213> Homo sapiens

<400> 32
Thr Pro Ala Ile Thr Arg Val Val Asp Phe Ala Lys Lys Leu Pro Met
1 5 10 15
Phe Ser Glu Leu Pro Cys Glu Asp Gln Ile Ile Leu Leu Lys Gly Cys
20 25 30

Cys

<210> 33

<211> 12

<212> PRT

<213> Homo sapiens

<400> 33

Leu Phe Pro Pro Leu Phe Leu Glu Val Phe Glu Asp
1 5 10

<210> 34

<211> 33

<212> PRT

<213> Homo sapiens

<400> 34

Thr Lys Cys Ile Ile Lys Ile Val Glu Phe Ala Lys Arg Leu Pro Gly
1 5 10 15

Phe Thr Gly Leu Ser Ile Ala Asp Gln Ile Thr Leu Leu Lys Ala Ala
20 25 30

Cys

<210> 35

<211> 12

<212> PRT

<213> Homo sapiens

<400> 35

Leu Phe Pro Pro Leu Phe Leu Glu Val Phe Glu Asp
1 5 10

<210> 36

<211> 33

<212> PRT

<213> Homo sapiens

<400> 36

Asp	Lys	Gln	Leu	Phe	Thr	Leu	Val	Glu	Trp	Ala	Lys	Arg	Ile	Pro	His
1				5					10					15	
Phe	Ser	Glu	Leu	Pro	Leu	Asp	Asp	Gln	Val	Ile	Leu	Leu	Lys	Ala	Gly
			20					25					30		

Trp

<210> 37

<211> 12

<212> PRT

<213> Homo sapiens

<400> 37

Pro	Ile	Asp	Thr	Phe	Leu	Met	Glu	Met	Leu	Glu	Ala
1				5					10		

<210> 38

<211> 33

<212> PRT

<213> Homo sapiens

<400> 38

Val	Glu	Ala	Val	Gln	Glu	Ile	Thr	Glu	Tyr	Ala	Lys	Asn	Ile	Pro	Gly
1				5					10					15	
Phe	Ile	Asn	Leu	Asp	Leu	Asn	Asp	Gln	Val	Thr	Leu	Leu	Lys	Tyr	Gly
			20					25					30		

Val

<210> 39

<211> 12

<212> PRT

<213> Homo sapiens

<400> 39

Ser	Leu	His	Pro	Leu	Leu	Gln	Glu	Ile	Tyr	Lys	Asp
1				5					10		

<210> 40

<211> 33

<212> PRT

<213> Homo sapiens

<400> 40

Ser	Tyr	Ser	Ile	Gln	Lys	Val	Ile	Gly	Phe	Ala	Lys	Met	Ile	Pro	Gly
1				5				10					15		
Phe	Arg	Asp	Leu	Thr	Ser	Glu	Asp	Gln	Ile	Val	Leu	Leu	Lys	Ser	Ser
			20					25					30		

Ala

<210> 41

<211> 12

<212> PRT

<213> Homo sapiens

<400> 41

Lys	Leu	Thr	Pro	Leu	Val	Leu	Glu	Val	Phe	Gly	Asn
1				5				10			

<210> 42

<211> 33

<212> PRT

<213> Homo sapiens

<220>

<221> MUTAGEN

<222> (12)..(12)

<223> Lys --> Ala (K362A)

<220>

<221> MUTAGEN

<222> (26)..(26)

<223> Val -->Arg (V376R)

<400> 42

Asp	Arg	Glu	Leu	Val	His	Met	Ile	Asn	Trp	Ala	Lys	Arg	Val	Pro	Gly
1				5				10					15		

Phe Val Asp Leu Thr Leu His Asp Gln Val His Leu Leu Glu Cys Ala
20 25 30

Trp

<210> 43

<211> 12

<212> PRT

<213> Homo sapiens

<220>

<221> MUTAGEN

<222> (8)..(8)

<223> Glu -->Lys (E542K)

<400> 43

Pro Leu Tyr Asp Leu Leu Leu Glu Met Leu Asp Ala
1 5 10

<210> 44

<211> 33

<212> PRT

<213> Homo sapiens

<400> 44

Gly Arg Gln Val Ile Ala Ala Val Lys Trp Ala Lys Ala Ile Pro Gly
1 5 10 15

Phe Arg Asn Leu His Leu Asp Asp Gln Met Thr Leu Leu Gln Tyr Ser
20 25 30

Trp

<210> 45

<211> 12

<212> PRT

<213> Homo sapiens

<400> 45

Glu Phe Pro Glu Met Leu Ala Glu Ile Ile Thr Asn
1 5 10

<210> 46

<211> 33

<212> PRT

<213> Homo sapiens

<400> 46

Glu Arg Gln Leu Leu Ser Val Val Lys Trp Ser Lys Ser Leu Pro Gly
1 5 10 15

Phe Arg Asn Leu His Ile Asp Asp Gln Ile Thr Leu Ile Gln Tyr Ser
20 25 30

Trp

<210> 47

<211> 12

<212> PRT

<213> Homo sapiens

<400> 47

Glu Phe Pro Glu Met Met Ser Glu Val Ile Ala Ala
1 5 10

<210> 48

<211> 33

<212> PRT

<213> Homo sapiens

<400> 48

Gly Lys Gln Met Ile Gln Val Val Lys Trp Ala Lys Val Leu Pro Gly
1 5 10 15

Phe Lys Asn Leu Pro Leu Glu Asp Gln Ile Thr Leu Ile Gln Tyr Ser
20 25 30

Trp

<210> 49

<211> 12

<212> PRT

<213> Homo sapiens

<400> 49

Glu Phe Pro Ala Met Leu Val Glu Ile Ile Ser Asp
1 5 10

<210> 50

<211> 33

<212> PRT

<213> Homo sapiens

<400> 50

Glu Arg Gln Leu Val His Val Val Lys Trp Ala Lys Ala Leu Pro Gly
1 5 10 15

Phe Arg Asn Leu His Val Asp Asp Gln Met Ala Val Ile Gln Tyr Ser
20 25 30

Trp

<210> 51

<211> 12

<212> PRT

<213> Homo sapiens

<400> 51

Asp Phe Pro Glu Met Met Ala Glu Ile Ile Ser Val
1 5 10

<210> 52

<211> 251

<212> PRT

<213> Homo sapiens

<220>

<221> VARIANT

<222> (1)..(251)

<223> Xaa is any amino acid

<220>

<223> Position 1 corresponds to position 211 of mature peptide

<400> 52

Lys	Pro	Glu	Pro	Thr	Asp	Glu	Glu	Trp	Glu	Leu	Ile	Lys	Thr	Val	Thr	
1				5					10					15		
Ala	Ala	His	Val	Ala	Thr	Asn	Ala	Gln	Gly	Ser	His	Trp	Lys	Asn	Lys	
			20					25					30			
Arg	Lys	Phe	Leu	Pro	Glu	Asp	Ile	Gly	Gln	Ala	Pro	Xaa	Xaa	Xaa	Xaa	
		35					40					45				
Xaa	Xaa	Gly	Gly	Lys	Val	Asp	Leu	Glu	Ala	Phe	Ser	His	Phe	Thr	Lys	
	50					55					60					
Ile	Ile	Thr	Pro	Ala	Ile	Thr	Arg	Val	Val	Asp	Phe	Ala	Lys	Lys	Leu	
65					70					75					80	
Pro	Met	Phe	Cys	Glu	Leu	Pro	Cys	Glu	Asp	Gln	Ile	Ile	Leu	Leu	Lys	
				85					90					95		
Gly	Cys	Cys	Met	Glu	Ile	Met	Ser	Leu	Arg	Ala	Ala	Val	Arg	Tyr	Asp	
			100					105					110			
Pro	Glu	Ser	Glu	Thr	Leu	Thr	Leu	Asn	Gly	Glu	Met	Ala	Val	Thr	Arg	
		115					120					125				
Gly	Gln	Leu	Lys	Asn	Gly	Gly	Leu	Gly	Val	Val	Ser	Asp	Ala	Ile	Phe	
	130					135					140					
Asp	Leu	Gly	Met	Ser	Leu	Ser	Ser	Phe	Asn	Leu	Asp	Asp	Thr	Glu	Val	
145					150					155					160	
Ala	Leu	Leu	Gln	Ala	Val	Leu	Leu	Met	Ser	Ser	Asp	Arg	Pro	Gly	Leu	
				165					170					175		
Ala	Cys	Val	Ala	Arg	Ile	Glu	Lys	Tyr	Gln	Asp	Ser	Phe	Leu	Leu	Ala	
		180						185					190			
Phe	Glu	His	Tyr	Ile	Asn	Tyr	Arg	Lys	His	His	Val	Thr	His	Phe	Trp	
		195					200					205				
Pro	Lys	Leu	Leu	Met	Lys	Val	Thr	Asp	Leu	Arg	Met	Ile	Gly	Ala	Cys	
	210					215					220					
His	Ala	Ser	Arg	Phe	Leu	His	Met	Lys	Val	Glu	Cys	Pro	Thr	Glu	Leu	
225					230					235					240	
Phe	Pro	Pro	Leu	Phe	Leu	Glu	Val	Phe	Glu	Asp						
				245					250							

<210> 53

<211> 250

<212> PRT

<213> Homo sapiens

<220>

<221> VARIANT

<222> (1)..(250)

<223> Xaa is any amino acid

<220>

<223> Position 1 corresponds to position 211 of mature peptide

<400> 53

Lys Pro Glu Pro Thr Asp Glu Glu Trp Glu Leu Ile Lys Thr Val Thr
1 5 10 15

Ala Ala His Val Ala Thr Asn Ala Gln Gly Ser His Trp Lys Asn Lys
20 25 30

Arg Lys Phe Leu Pro Glu Asp Ile Gly Gln Ala Pro Xaa Xaa Xaa Xaa
35 40 45

Xaa Xaa Gly Gly Lys Val Asp Leu Glu Ala Phe Ser His Phe Thr Lys
50 55 60

Ile Ile Thr Pro Ala Ile Thr Arg Val Val Asp Phe Ala Lys Lys Leu
65 70 75 80

Pro Met Phe Cys Glu Leu Pro Cys Glu Asp Gln Ile Ile Leu Leu Lys
85 90 95

Gly Cys Cys Met Glu Ile Met Ser Leu Arg Ala Ala Val Arg Tyr Asp
100 105 110

Pro Glu Ser Glu Thr Leu Thr Leu Asn Gly Glu Met Ala Val Thr Arg
115 120 125

Gly Gln Leu Lys Asn Gly Leu Gly Val Val Ser Asp Ala Ile Phe Asp
130 135 140

Leu Gly Met Ser Leu Ser Ser Phe Asn Leu Asp Asp Thr Glu Val Ala
145 150 155 160

Leu Leu Gln Ala Val Leu Leu Met Ser Ser Asp Arg Pro Gly Leu Ala
165 170 175

Cys Val Ala Arg Ile Glu Lys Tyr Gln Asp Ser Phe Leu Leu Ala Phe
180 185 190

Glu His Tyr Ile Asn Tyr Arg Lys His His Val Thr His Phe Trp Pro
195 200 205

Lys Leu Leu Met Lys Val Thr Asp Leu Arg Met Ile Gly Ala Cys His
210 215 220

Ala Ser Arg Phe Leu His Met Lys Val Glu Cys Pro Thr Glu Leu Phe
225 230 235 240

Pro Pro Leu Phe Leu Glu Val Phe Glu Asp
245 250

<210> 54

<211> 13

<212> PRT

<213> Homo sapiens

<220>

<223> Position 1 corresponds to position 686 of mature peptide

<400> 54

Lys His Lys Ile Leu His Arg Leu Leu Gln Asp Ser Ser
1 5 10

<210> 55

<211> 9

<212> PRT

<213> Homo sapiens

<220>

<223> Position 1 corresponds to position 688 of mature peptide

<400> 55

Lys Ile Leu His Arg Leu Leu Gln Asp
1 5

<210> 56

<211> 245

<212> PRT

<213> Homo sapiens

<220>

<223> Position 1 corresponds to position 305 of mature peptide

<400> 56

Ser Leu Ala Leu Ser Leu Thr Ala Asp Gln Met Val Ser Ala Leu Leu
1 5 10 15

Asp Ala Glu Pro Pro Ile Leu Tyr Ser Glu Tyr Asp Pro Thr Arg Pro
20 25 30

Phe Ser Glu Ala Ser Met Met Gly Leu Leu Thr Asn Leu Ala Asp Arg
35 40 45

Glu Leu Val His Met Ile Asn Trp Ala Lys Arg Val Pro Gly Phe Val
50 55 60

Asp Leu Thr Leu His Asp Gln Val His Leu Leu Glu Cys Ala Trp Leu
65 70 75 80

Glu Ile Leu Met Ile Gly Leu Val Trp Arg Ser Met Glu His Pro Gly
85 90 95

Lys	Leu	Leu	Phe	Ala	Pro	Asn	Leu	Leu	Leu	Asp	Arg	Asn	Gln	Gly	Lys
			100					105					110		
Cys	Val	Glu	Gly	Met	Val	Glu	Ile	Phe	Asp	Met	Leu	Leu	Ala	Thr	Ser
		115					120					125			
Ser	Arg	Phe	Arg	Met	Met	Asn	Leu	Gln	Gly	Glu	Glu	Phe	Val	Cys	Leu
	130					135					140				
Lys	Ser	Ile	Ile	Leu	Leu	Asn	Ser	Gly	Val	Tyr	Thr	Phe	Leu	Ser	Ser
145					150					155					160
Thr	Leu	Lys	Ser	Leu	Glu	Glu	Lys	Asp	His	Ile	His	Arg	Val	Leu	Asp
				165					170					175	
Lys	Ile	Thr	Asp	Thr	Leu	Ile	His	Leu	Met	Ala	Lys	Ala	Gly	Leu	Thr
			180					185					190		
Leu	Gln	Gln	Gln	His	Gln	Arg	Leu	Ala	Gln	Leu	Leu	Leu	Ile	Leu	Ser
		195					200					205			
His	Ile	Arg	His	Met	Ser	Asn	Lys	Gly	Met	Glu	His	Leu	Tyr	Ser	Met
	210					215					220				
Lys	Cys	Lys	Asn	Val	Val	Pro	Leu	Tyr	Asp	Leu	Leu	Leu	Glu	Met	Leu
225				230						235					240
Asp	Ala	His	Arg	Leu											
				245											

<210> 57

<211> 237

<212> PRT

<213> Homo sapiens

<220>

<223> Position 1 corresponds to position 305 of mature peptide

<400> 57

Ser	Leu	Ala	Leu	Ser	Leu	Thr	Ala	Asp	Gln	Met	Val	Ser	Ala	Leu	Leu
1				5					10					15	
Asp	Ala	Glu	Pro	Pro	Ile	Leu	Tyr	Ser	Glu	Tyr	Asp	Pro	Thr	Arg	Pro
			20					25					30		
Phe	Ser	Glu	Ala	Ser	Met	Met	Gly	Leu	Leu	Thr	Asn	Leu	Ala	Asp	Arg
		35					40					45			
Glu	Leu	Val	His	Met	Ile	Asn	Trp	Ala	Lys	Lys	Arg	Val	Pro	Gly	Phe
	50					55					60				
Val	Asp	Leu	Thr	Leu	His	Asp	Gln	Val	His	Leu	Leu	Glu	Cys	Ala	Trp
65					70					75					80
Leu	Glu	Ile	Leu	Met	Ile	Gly	Leu	Val	Trp	Arg	Ser	Met	Glu	His	Pro
				85				90						95	
Gly	Lys	Leu	Leu	Phe	Ala	Pro	Asn	Leu	Leu	Leu	Asp	Arg	Asn	Gln	Gly
			100					105						110	

Lys Cys Val Gly Gly Met Val Glu Ile Phe Asp Met Leu Leu Ala Thr
 115 120 125
 Ser Ser Arg Phe Arg Met Met Asn Leu Gln Gly Glu Glu Phe Val Cys
 130 135 140
 Leu Lys Ser Ile Ile Leu Leu Asn Ser Gly Val Tyr Thr Phe Glu Lys
 145 150 155 160
 Asp His Ile His Arg Val Leu Asp Lys Ile Thr Asp Thr Leu Ile His
 165 170 175
 Leu Met Ala Lys Ala Gly Leu Thr Leu Gln Gln Gln His Gln Arg Leu
 180 185 190
 Ala Gln Leu Leu Leu Ile Leu Ser His Ile Arg His Met Ser Asn Lys
 195 200 205
 Gly Met Glu His Leu Tyr Ser Met Lys Cys Lys Asn Val Val Pro Leu
 210 215 220
 Tyr Asp Leu Leu Leu Glu Met Leu Asp Ala His Arg Leu
 225 230 235

<210> 58

<211> 11

<212> PRT

<213> Homo sapiens

<220>

<223> Position 1 corresponds to position 687 of mature peptide

<400> 58

His Lys Ile Leu His Arg Leu Leu Gln Asp Ser
 1 5 10

<210> 59

<211> 246

<212> PRT

<213> Homo sapiens

<220>

<223> Position 1 corresponds to position 306 of mature peptide

<400> 59

Leu Ala Leu Ser Leu Thr Ala Asp Gln Met Val Ser Ala Leu Leu Asp
 1 5 10 15

Ala Glu Pro Pro Ile Leu Tyr Ser Glu Tyr Asp Pro Thr Arg Pro Phe
 20 25 30

Ser	Glu	Ala	Ser	Met	Met	Gly	Leu	Leu	Thr	Asn	Leu	Ala	Asp	Arg	Glu
		35					40					45			
Leu	Val	His	Met	Ile	Asn	Trp	Ala	Lys	Arg	Val	Pro	Gly	Phe	Val	Asp
	50					55					60				
Leu	Thr	Leu	His	Asp	Gln	Val	His	Leu	Leu	Glu	Cys	Ala	Trp	Leu	Glu
65					70					75					80
Ile	Leu	Met	Ile	Gly	Leu	Val	Trp	Arg	Ser	Met	Glu	His	Pro	Gly	Lys
				85					90					95	
Leu	Leu	Phe	Ala	Pro	Asn	Leu	Leu	Leu	Asp	Arg	Asn	Gln	Gly	Lys	Cys
			100					105					110		
Val	Glu	Gly	Met	Val	Glu	Ile	Phe	Asp	Met	Leu	Leu	Ala	Thr	Ser	Ser
		115					120					125			
Arg	Phe	Arg	Met	Met	Asn	Leu	Gln	Gly	Glu	Glu	Phe	Val	Cys	Leu	Lys
	130					135					140				
Ser	Ile	Ile	Leu	Leu	Asn	Ser	Gly	Val	Tyr	Thr	Phe	Leu	Ser	Ser	Thr
145					150					155					160
Leu	Lys	Ser	Leu	Glu	Glu	Lys	Asp	His	Ile	His	Arg	Val	Leu	Asp	Lys
			165						170					175	
Ile	Thr	Asp	Thr	Leu	Ile	His	Leu	Met	Ala	Lys	Ala	Gly	Leu	Thr	Leu
			180					185						190	
Gln	Gln	Gln	His	Gln	Arg	Leu	Ala	Gln	Leu	Leu	Leu	Ile	Leu	Ser	His
		195					200					205			
Ile	Arg	His	Met	Ser	Asn	Lys	Gly	Met	Glu	His	Leu	Tyr	Ser	Met	Lys
	210					215					220				
Cys	Lys	Asn	Val	Val	Pro	Leu	Tyr	Asp	Leu	Leu	Leu	Glu	Met	Leu	Asp
225					230					235					240
Ala	His	Arg	Leu	His	Ala										
				245											

<210> 60

<211> 11

<212> PRT

<213> Homo sapiens

<220>

<223> Position 1 corresponds to position 686 of mature peptide

<400> 60

Lys	His	Lys	Ile	Leu	His	Arg	Leu	Leu	Gln	Asp
1				5					10	